### **ATP-III Treat to Goal Statin Dosing Guide**

(from baseline LDL-C)

The primary focus of treatment for the prevention of coronary heart disease (CHD) is the reduction of LDL-C. Treatment intensity is stratified by risk into 3 groups as defined by the National Cholesterol Education Program's Adult Treatment Panel—III (ATP-III) guidelines published in May 2001. Patients will fall into one of three groups with a corresponding LDL goal as defined below based on a CHD risk assessment.

| Blood Level | Treatment Groups |           |           |
|-------------|------------------|-----------|-----------|
| LDL-C       | Group I          | Group II  | Group III |
| (mg/dL)     | LDL Goal:        | LDL Goal: | LDL Goal: |
|             | 160              | 130       | 100       |
| 110         |                  |           |           |
| 115         |                  |           |           |
| 120         |                  | At Goal   | S-5       |
| 125         |                  |           |           |
| 130         |                  |           |           |
| 135         | At Goal          |           | S-10      |
| 140         |                  |           | 0.10      |
| 145         |                  |           |           |
| 150         |                  | S-5       | S-20      |
| 155         |                  |           |           |
| 160         |                  |           | S-40      |
| 165         |                  |           |           |
| 170         |                  |           |           |
| 175<br>180  | S-5              | S-10      |           |
| 185         |                  |           | S-80      |
| 190         |                  |           |           |
| 195         |                  | S-20      |           |
| 200         |                  | 0-20      |           |
| 210         | S-10             | S-40      | A-40*     |
| 220         |                  | 0 40      |           |
| 230         |                  |           | A-80*     |
| 240         | S-20             | S-80      |           |
| 250         | S-40             |           |           |
| 260         | 3-40             | A-40*     |           |
| 270         |                  |           |           |
| 280         | S-80             |           |           |
| 290         |                  | A-80*     |           |
| 300         |                  |           |           |
| 310         |                  |           |           |
| 320         | A-40*            |           |           |
| 330         | A-80*            |           |           |
| 340         |                  |           |           |
| 350         |                  |           |           |

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|-----------------------|--|--|
| Group I               | This group consists of primary prevention patients with low risk for atherosclerosis. Low risk is defined as non-diabetics with no or one risk factors and a <10% 10-year CHD risk as determined by their Framingham Score. The LDL goal is 160mg/dL.  |  |
| Group II              | This group consists of primary prevention patients with moderate risk for atherosclerosis. Moderate risk is defined as non-diabetics with two or more CHD risk factors and a 10-20% 10-year CHD risk as determined by their Framingham Score. The LDL goal is 130mg/dL.  |  |
| Group III             | This group consists of patients with present CHD, diabetics and other CHD risk equivalent patients. CHD risk eqivalents include patients with diabetes, peripheral arterial disease, symptomactic carotid artery disease, abdominal aortic aneurysm or a >20% 10-year CHD risk as determined by their Framingham Score. The LDL goal is <100mg/dL. |  |
| S-5                   | Simvastatin 5mg  |  |
| S-10                  | Simvastatin 10mg   |  |
| S-20                  | Smvastatin 20mg  |  |
| S-40                  | Simvastatin 40mg   |  |
| S-80                  | Simvastatin 80mg   |  |
| A-40*                 | Atorvastatin 40mg *Special Order Req.*   |  |
| A-80*                 | Atorvastatin 80mg *Special Order Req.*   |  |
|                       | Beyond Documented Limits   |  |

#### **CHD Risk Factors:**

Age -- Male ≥ 45

Female ≥ 55, or premature menopause w/o estrogen replacement

Family History (definite MI or sudden death at <55 in father/1st degree relative;<65 mother/1st degree relative) Cigarette smoking, including ex-smokers <1 yr

Hypertension (≥ 140/90mmHg-on more than one occasion, or on antihypertensive medication)

Low HDL <40mg/dL

Negative Risk Factor: (subtract one risk factor)

High HDL ≥ 60mg/dL

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| 145         |           |                |           |
| 150         |           | S-5            | S-20      |
| 155         |           |                |           |
| 160         |           |                | S-40      |
| 165         | S-5       |                |           |
| 170         |           |                |           |
| 175         |           | S-10           |           |
| 180         |           |                | S-80      |
| 185         |           |                |           |
| 190         |           |                |           |
| 195         |           | S-20           |           |
| 200         |           | 2 12           | A-40*     |
| 210         | 0.15      | S-40           |           |
| 220         | S-10      |                | A 00*     |
| 230         | 2.00      | S-80           | A-80*     |
| 240<br>250  | S-20      |                |           |
| 250<br>260  | S-40      | A-40*          |           |
| 270         |           | A-40           |           |
| 280         |           |                |           |
| 290         | S-80      | A-80*          |           |
| 300         | <u> </u>  |                |           |
| 310         |           |                |           |
| 320         | A-40*     |                |           |
| 330         | 71 40     |                |           |
| 340         | A-80*     |                |           |
| 350         |           |                |           |

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| S-5<br>S-10<br>S-20<br>S-40<br>S-80<br>A-40*<br>A-80* | Simvastatin 5mg Simvastatin 10mg Smvastatin 20mg Simvastatin 40mg Simvastatin 80mg Atorvastatin 40mg *Special Order Req.* Atorvastatin 80mg *Special Order Req.* Beyond Documented Limits   |  |

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### **Prescribing Notes**

| Dose QPM (atorvastatin can be QAM)  |
|---|
| BID dosing less effective than 2X QPM   |
| No evidence that titration is necessary, but common with Sim 80, Ator 40&80                                   |
| General guide, adapt to patient-specific concerns   |
| Lipid Profile: Baseline, ≥ 6wks after start or dose change, yearly after achieving goal                       |
| LFTs: Baseline, ≥ 4wks of high dose statin (Sim 80mg, Ator 40&80mg), Q 6-12mo as needed                       |
| CPKs: not routine, only if complications occur  |
| Statin use is ALWAYS in addition to lifestyle modifications, may require reminders to pts                     |
| Impact varies by patient, usually a 5-15% reduction in LDL levels   |
| Patients can eat through the best statins available   |
| Statins are not first line agents (use niacin or gemfibrozil, ↑ HDL 20-35%)                                   |
| All statin ↑ HDL levels, but not usually more than 10%  |
| Statins are not first line agents (use niacin or gemfibrozil, ↓ TG 30-60%)                                    |
| Greater TG baseline yields greater statin response, limited effect at high (200-500) TG levels                |
| Statin+ fibrate or niacin increases risk of rhabdomyolysis  |
| All statins have warnings in their labeling stating increased risk of myopathies when combined with fibrates  |
| Simvastatin 10mg is the only statin labeled for concomitant use with fibrates                                 |
| Simvastatin, atorvastatin, & lovastatin are metabolized via hepatic CYP 450 3A4 - avoid 3A4 inhibitors        |
| Pravastatin is renally excreted (Note: max LDL reduction is ~37% with 80mg) and free of 3A4 drug interactions |
| Common 3A4 inhibitors are listed in the table below (Non-Formulary Requests Section)                          |
|   |

## **Non-Formulary Requests**

MTFs are required to have simvastatin as the only formulary statin. When simvastatin does not meet the clinical needs of the patient and medical necessity is established, MTFs should obtain the most appropriate alternative. Wherever possible, procedures should be streamlined for these cases, requiring little, if any, involvement by the patient in the process.

Please contact the Pharmacy Department for your MTF-specific procedures.

Examples of medical necessity include:

| Potential<br>Drug<br>Interactions | Chronic meds: amiodarone, cyclosporine, HIV protease inhibitors, nefazodone, -azole antifungals [itraconazole, ketoconazole & fluconazole (if dose >200mg/day)], & macolides (erythro-, clarithro-, only in chronic use, else see Short Term meds)  Short Term meds: DO NOT CONVERT, temporary cessation of simvastatin most prudent |
|-----------------------------------|--|
| Inadequate<br>Therapy             | Patients not achieving their LDL goal with simvastatin 80mg or new patients not expected to achieve their LDL goal with simvastatin 80mg (based on LDL results and chart provided) should begin atorvastatin (40mg or 80mg as required to attain goal) immediately.  |
| Adverse<br>Reactions              | Beyond common initial self-limiting events (i.e., gastrointestinal distress)   |
| Allergy                           | History of simvastatin allergy   |

